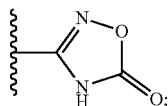
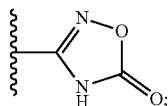


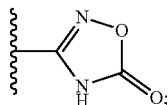
K is C(O)NH<sub>2</sub>, COOH, SO<sub>3</sub>H, OSO<sub>3</sub>H, PO<sub>3</sub>H<sub>2</sub>, OPO<sub>3</sub>H<sub>2</sub>, NH<sub>2</sub>, NHR<sub>19</sub>, NR<sub>19</sub>R<sub>20</sub>, SO<sub>2</sub>R<sub>21</sub>, glycoside, lower C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, C<sub>6</sub> alkoxy, or



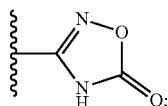
L is aryl, OH, C(O)NH<sub>2</sub>, COOH, SO<sub>3</sub>H, OSO<sub>3</sub>H, PO<sub>3</sub>H<sub>2</sub>, OPO<sub>3</sub>H<sub>2</sub>, NH<sub>2</sub>, NHR<sub>19</sub>, NR<sub>19</sub>R<sub>20</sub>, SO<sub>2</sub>R<sub>21</sub>, glycoside, lower C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, C<sub>6</sub> alkoxy, or



M is aryl, OH, C(O)NH<sub>2</sub>, COOH, SO<sub>3</sub>H, OSO<sub>3</sub>H, PO<sub>3</sub>H<sub>2</sub>, OPO<sub>3</sub>H<sub>2</sub>, NH<sub>2</sub>, NHR<sub>19</sub>, NR<sub>19</sub>R<sub>20</sub>, SO<sub>2</sub>R<sub>21</sub>, glycoside, lower C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, C<sub>6</sub> alkoxy, or



Q is aryl, OH, C(O)NH<sub>2</sub>, COOH, SO<sub>3</sub>H, OSO<sub>3</sub>H, PO<sub>3</sub>H<sub>2</sub>, OPO<sub>3</sub>H<sub>2</sub>, NH<sub>2</sub>, NHR<sub>19</sub>, NR<sub>19</sub>R<sub>20</sub>, SO<sub>2</sub>R<sub>21</sub>, glycoside, lower C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, C<sub>6</sub> alkoxy, or



R<sub>19</sub>, R<sub>20</sub> and R<sub>21</sub> are independently C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, or C<sub>6</sub> alkyl or R<sub>19</sub> and R<sub>20</sub> taken together with the attached nitrogen atom form a five membered ring;

V is a bond, —CH<sub>2</sub>—, —CH<sub>2</sub>CH<sub>2</sub>—, —CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>—, —O—CH<sub>2</sub>—, —OCH<sub>2</sub>CH<sub>2</sub>— or —OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>—;

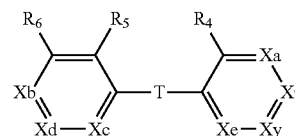
R<sub>12</sub>, R<sub>13</sub>, R<sub>14</sub>, R<sub>15</sub>, R<sub>16</sub>, R<sub>n</sub>, and R<sub>18</sub>, are, independently, H or C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, or C<sub>6</sub> alkyl; and

Z is (CHR<sub>1</sub>)<sub>n</sub>—C(O)—NR<sub>2</sub>(CHR<sub>3</sub>)<sub>m</sub>—Ar, where Ar is a substituted or unsubstituted aryl or nitrogen-containing heteroaryl group, R<sub>1</sub>, R<sub>2</sub>, and R<sub>3</sub> are independently H or C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, or C<sub>6</sub> alkyl; and

n and m are, independently 0, 1, or 2;

provided that at least one of R<sub>a</sub>, R<sub>b</sub>, R<sub>c</sub>, R<sub>d</sub>, R<sub>e</sub>, R<sub>4</sub>, R<sub>5</sub>, and R<sub>6</sub> is P.

2. A method of protecting against or treating hearing loss or osteoporosis in a subject comprising administering a compound having the Formula I:



(Formula I)

or a salt, solvate, hydrate, or prodrug thereof, wherein:

T is absent, CR<sub>12</sub>R<sub>13</sub>, C(O), O, S, S(O), S(O)<sub>2</sub>, NR<sub>14</sub>, C(R<sub>15</sub>R<sub>16</sub>)C(R<sub>17</sub>R<sub>18</sub>), CH<sub>2</sub>O, or OCH<sub>2</sub>;

X<sub>y</sub> is CZ, CY, N, or N—O;

X<sub>z</sub> is CZ, CY, N, or N—O;

at least one of X<sub>y</sub> and X<sub>z</sub> is CZ;

Y is selected from hydrogen, hydroxyl, halogen, lower (C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, or C<sub>6</sub>) alkyl, C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, or C<sub>6</sub> alkoxy, O-lower (C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, or C<sub>6</sub>) alkyl-aryl, and O-benzyl;

X<sub>a</sub> is CR<sub>a</sub> or N, or N—O;

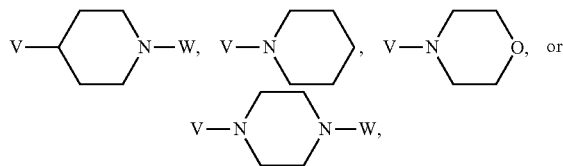
X<sub>b</sub> is CR<sub>b</sub>, N, or N—O;

X<sub>c</sub> is CR<sub>c</sub> or N, or N—O;

X<sub>d</sub> is CR<sub>d</sub> or N, or N—O;

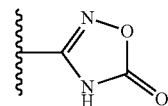
X<sub>e</sub> is CR<sub>e</sub>, N, or N—O;

R<sub>a</sub>, R<sub>b</sub>, R<sub>c</sub>, R<sub>d</sub>, R<sub>e</sub>, R<sub>4</sub>, R<sub>5</sub>, and R<sub>6</sub> are, independently, hydrogen, hydroxyl, halogen, P, C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, or C<sub>6</sub> alkyl, C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, or C<sub>6</sub> alkoxy, O-lower (C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, or C<sub>6</sub>) alkyl-aryl, O-benzyl, C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, or C<sub>6</sub> alkyl-OH, COOH, COO-lower (C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, or C<sub>6</sub>) alkyl, SO<sub>2</sub>H, SO<sub>2</sub>-lower (C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, or C<sub>6</sub>) alkyl,



where W is H, or C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, or C<sub>6</sub> alkyl, C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, or C<sub>6</sub> alkyl-aryl;

P is SO<sub>3</sub>H, OSO<sub>3</sub>H, OPO<sub>3</sub>H<sub>2</sub>, OPO<sub>3</sub>H<sub>2</sub>, NH<sub>2</sub>, NHR<sub>19</sub>, NHR<sub>2</sub>OR<sub>21</sub>,



tetrazole, O-lower (C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, or C<sub>6</sub>) alkyl-K, O—C(O)-lower (C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, or C<sub>6</sub>) alkyl-L, NH-lower (C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, or C<sub>6</sub>) alkyl-M, or O-aryl-Q, further wherein lower alkyl is linear or branched alkyl;

K is C(O)NH<sub>2</sub>, COOH, SO<sub>3</sub>H, OSO<sub>3</sub>H, PO<sub>3</sub>H<sub>2</sub>, OPO<sub>3</sub>H<sub>2</sub>, NH<sub>2</sub>, NHR<sub>19</sub>, NR<sub>19</sub>R<sub>20</sub>, SO<sub>2</sub>R<sub>21</sub>, glycoside, lower C<sub>1</sub>, C<sub>2</sub>, C<sub>3</sub>, C<sub>4</sub>, C<sub>5</sub>, C<sub>6</sub> alkoxy, or